09/787436

(FILE 'REGISTRY' ENTERED AT 12:53:17 ON 22 NOV 2004) STR L13 17 19 NH 12 10 - CH-√ C NH ~ C ~ NH ~ G1~ 11 15 14. 13 16 NH-\rightarrow CH2-\rightarrow CH3 N 1 8 18

Str.

REP G1=(3-3) CH2
NODE ATTRIBUTES:
CONNECT IS X2 RC AT 4
CONNECT IS X2 RC AT 5
DEFAULT MLEVEL IS ATOM
DEFAULT ECLEVEL IS LIMITED

GRAPH ATTRIBUTES: RING(S) ARE ISOLATED OR EMBEDDED NUMBER OF NODES IS 19

STEREO ATTRIBUTES: NONE

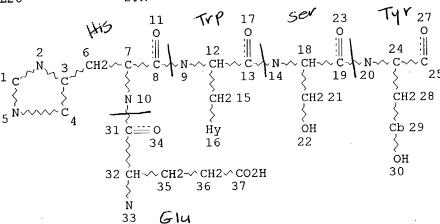
L15 L26

المراجية

16 E .

CANAL C

2838 SEA FILE=REGISTRY SSS FUL L13 STR



NODE ATTRIBUTES:
DEFAULT MLEVEL IS ATOM
GGCAT IS PCY AT 16
GGCAT IS MCY UNS AT 29
DEFAULT ECLEVEL IS LIMITED

GRAPH ATTRIBUTES:
RING(S) ARE ISOLATED OR EMBEDDED
NUMBER OF NODES IS 36

Searcher :

Shears

STEREO ATTRIBUTES: NONE

L27 3 SEA FILE=REGISTRY SUB=L15 SSS FUL L26

100.0% PROCESSED 74 ITERATIONS

3 ANSWERS

SEARCH TIME: 00.00.01

FILE 'CAPLUS' ENTERED AT 12:57:42 ON 22 NOV 2004

L28 · 3 S L27

L28 ANSWER 1 OF 3 CAPLUS COPYRIGHT 2004 ACS on STN

ACCESSION NUMBER:

1989:112150 CAPLUS

DOCUMENT NUMBER:

110:112150

TITLE:

754 C.

Luliberin peptide analog for stimulation of production

of roe during artificial milting of common tench (Tinca tinca), trout (Micropterus samonides) and

sheatfish (Silurus glanis)

INVENTOR(S):

Kouril, Jan; Hamackova, Jitka; Machacek, Jiri; Barth,

Tomislav; Flegel, Martin

PATENT ASSIGNEE(S):

: Czech.

SOURCE:

Czech., 8 pp.

CODEN: CZXXA9

DOCUMENT TYPE:

Patent

LANGUAGE:

Czech

FAMILY ACC. NUM. COUNT: 3

PATENT INFORMATION:

PATENT NO.	KIND	DATE	AP	PLICATION NO.	DATE
/ 					
CS 244397	B1	19860717	CS	1984-8441	19841006
PRIORITY APPLN. INFO.:			CS	1984-8441	19841006

AB A method of obtaining roe during artificial milting of T. tinca, M. samonides, and S. glanis is characterized by injecting female egg-carrying fish, during premilting maturity, i.m. with Glu-His-Trp-Ser-Tyr-D-Ala-Leu-Arg-Pro-NHEt (I), an analog of luliberin, at 5-20 μg/kg. Female egg-carrying tench (500-1000 g, 4-5 yr) at 20-22° were injected i.m. with I (5-80 μg/kg). Within 15 h, the roe were ready for milting. A dosage of 20 μg/kg resulted in ovulation by 62.5% of the fish. The most effective dosage range was 5-20 μg/kg.

IT 119261-01-7

RL: BIOL (Biological study)

(ovulation stimulation by, in fish)

RN 119261-01-7 CAPLUS

CN Luteinizing hormone-releasing factor (swine), 1-L-glutamic acid-6-D-alanine-9-(N-ethyl-L-prolinamide)-10-deglycinamide- (9CI) (CA INDEX NAME)

Absolute stereochemistry.

Searcher : Shears

PAGE 1-A

PAGE 1-B

=NH

\$25 P.

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2-x3

L28 ANSWER 2 OF 3 CAPLUS COPYRIGHT 2004 ACS on STN

ACCESSION NUMBER: DOCUMENT NUMBER:

1988:556330 CAPLUS

109:156330

TITLE:

Reversed-phase high-performance liquid chromatography

of fertirelin acetate and related compounds

AUTHOR(S):

Hartman, Patrick A.; Stodola, John D.

CORPORATE SOURCE:

Upjohn Co., Kalamazoo, MI, 49001, USA

SOURCE:

Journal of Chromatography (1988), 444, 177-82

CODEN: JOCRAM; ISSN: 0021-9673

Searcher :

Shears

DOCUMENT TYPE:

Journal

LANGUAGE:

English

AB Separation of fertirelin acetate (FA) from process impurities, potential degradation products and related peptides including LH releasing hormone was achieved by reversed-phase HPLC. A number of chromatog. conditions (column type mobile phase composition, isocratic/gradient elution) and detection systems were utilized to examine the bulk drug and formulation of FA. Examples of sepns. designed for potency and impurity detns. are described. Complete recovery of FA is obtained with an isocratic HPLC system. An external standard method is used to determine potency with a precision of

<18

75 O. L.

 $\ensuremath{\mathtt{R.S.D.}}$ A gradient HPLC system is used to determine impurities with a precision

of 5-10% R.S.D. at the 1-2% impurity level. As little as .apprx.0.1% (area %) of related peptides are detected at 214 mm.

IT 116921-32-5

RL: ANT (Analyte); ANST (Analytical study)
 (HPLC of, as fertirelin acetate impurity)

RN 116921-32-5 CAPLUS

CN Luteinizing hormone-releasing factor (swine), 1-L-glutamic acid-9-(N-ethyl-L-prolinamide)-10-deglycinamide- (9CI) (CA INDEX NAME)

Absolute stereochemistry.

PAGE 1-A

Searcher :

Shears

 $\approx_{\rm NH}$

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L28 ANSWER 3 OF 3 CAPLUS COPYRIGHT 2004 ACS on STN

ACCESSION NUMBER:

1988:82207 CAPLUS

DOCUMENT NUMBER:

108:82207

TITLE:

HPLC of leuprolide acetate in injectable solutions

AUTHOR(S): Sutherland, J. W.; Menon, G. N.

CORPORATE SOURCE:

Pharm. Prod. Div., Abbott Lab., North Chicago, IL,

60064, USA

SOURCE:

Journal of Liquid Chromatography (1987), 10(10),

2281-9

CODEN: JLCHD8; ISSN: 0148-3919

DOCUMENT TYPE:

Journal

LANGUAGE:

English

AB A stability-indicating HPLC method based on a $5-\mu$ octadecylsilane column and pH 6.5 0.057M aqueous monobasic ammonium phosphate solution-MeCN (77:23 by volume) mobile phase, and UV detection at 220 nm was used for the determination of leuprolide acetate in injections. Et p-hydroxybenzoate was the

internal standard The relative standard deviation was 1.8% and the method was

also successfully used for the determination of impurities/precursors occurring

during the drug manufacture. The drug was more stable at pH 3.3 than at pH 10.3

when heated at 100° for 16 h. Glul-leuprolide was the major degradation product when the drug was treated with 0.1N HCl at 40° for 48 h.

IT 112642-13-4

RL: ANT (Analyte); ANST (Analytical study) (determination of, as leuprolide impurity by HPLc)

RN 112642-13-4 CAPLUS

CN Luteinizing hormone-releasing factor (swine), 1-L-glutamic acid-6-D-leucine-9-(N-ethyl-L-prolinamide)-10-deglycinamide- (9CI) (CA INDEX NAME)

Absolute stereochemistry.

PAGE 1-A

PAGE 1-B

 \approx_{NH}

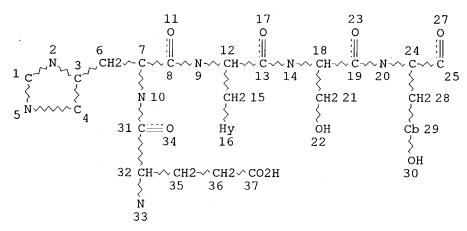
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FILE 'CAOLD' ENTERED AT 12:58:05 ON 22 NOV 2004 L29 0 S L27

FILE 'USPATFULL' ENTERED AT 12:58:10 ON 22 NOV 2004 L30 0 S L27

FILE 'MEDLINE, BIOSIS, EMBASE' ENTERED AT 12:58:19 ON 22 NOV 2004 L31 0 S L27

(FILE 'MARPAT' ENTERED AT 12:58:35 ON 22 NOV 2004) STR



NODE ATTRIBUTES:

L32

3524 L

DEFAULT MLEVEL IS ATOM

MLEVEL IS CLASS AT 16 29

GGCAT IS PCY AT 16

GGCAT IS MCY UNS AT 29

DEFAULT ECLEVEL IS LIMITED

GRAPH ATTRIBUTES:

RING(S) ARE ISOLATED OR EMBEDDED

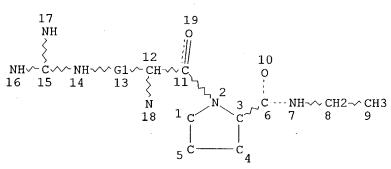
NUMBER OF NODES IS 36

STEREO ATTRIBUTES: NONE

ATTRIBUTES SPECIFIED AT SEARCH-TIME: ECLEVEL IS LIM ON ALL NODES

ALL RING(S) ARE ISOLATED

L34 92 SEA FILE=MARPAT SSS FUL L32 (MODIFIED ATTRIBUTES) L35 STR



REP G1=(3-3) CH2 NODE ATTRIBUTES:

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CONNECT IS X2 RC AT

CONNECT IS X2 RC AT

Searcher :

Shears

DEFAULT MLEVEL IS ATOM DEFAULT ECLEVEL IS LIMITED

GRAPH ATTRIBUTES:

راج ويعير

RING(S) ARE ISOLATED OR EMBEDDED

NUMBER OF NODES IS 19

STEREO ATTRIBUTES: NONE

ATTRIBUTES SPECIFIED AT SEARCH-TIME:

ECLEVEL IS LIM ON ALL NODES ALL RING(S) ARE ISOLATED

L36 6 SEA FILE=MARPAT SUB=L34 SSS FUL L35 (MODIFIED ATTRIBUTES)

100.0% PROCESSED 82 ITERATIONS 6 ANSWERS

SEARCH TIME: 00.00.01

L36 ANSWER 1 OF 6 MARPAT COPYRIGHT 2004 ACS on STN

ACCESSION NUMBER:

141:162339 MARPAT

TITLE:

Improved linkers for pharmaceutical compounds

INVENTOR(S):

De Haeen, Christoph; Nunn, Adrian; Swenson, Rolf E.

PATENT ASSIGNEE(S):

Bracco Imaging S.P.A., Italy

SOURCE:

PCT Int. Appl., 122 pp.

CODEN: PIXXD2

DOCUMENT TYPE:

Patent

LANGUAGE:

English

FAMILY ACC. NUM. COUNT:

PATENT INFORMATION:

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PATENT NO.
                                          KIND DATE
                                                                                                     APPLICATION NO.
                                                                                                                                                   DATE
----- · ----
                                                           -----
                                                                                                     _____
WO 2004062574
                                            A2
                                                             20040729
                                                                                                 WO 2003-US41656 20031224
          W: AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, CA, CH, CN, CU, CZ, DE,
         W: AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, CA, CH, CN, CU, CZ, DE, DK, EE, ES, FI, GB, GD, GE, GH, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MD, MG, MK, MN, MW, MX, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SK, SL, TJ, TM, TR, TT, UA, UG, US, UZ, VN, YU, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM

RW: BW, GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZM, ZW, AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, HU, IE, IT, LU, MC, NL, PT, RO, SE, SI, SK, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG

APPLN: INFO::

US 2003-439722 P. 20030112
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PRIORITY APPLN: INFO.:

US 2003-439722P 20030113

A new and improved method for extending the half life of pharmaceutical compds. for use in diagnostic imaging or therapy uses a novel linker to attach a diagnostic or therapeutic moiety to a targeting peptide or another diagnostic or therapeutic moiety. The resulting compound may have the general formula M-N-O-P-Q, wherein M is the diagnostic or therapeutic moiety, N-O-P is the linker of the present invention, and Q is the targeting peptide. In another embodiment the compds. may have the formula M-N-O-P-M, wherein M is independently a diagnostic or therapeutic moiety and N-O-P is the linker of the invention. Methods for imaging or treating a patient using the compds. of the invention are also provided. Methods and kits for preparing a diagnostic imaging agent from the compound are

further

provided. Methods for radiotherapy of a patient using the compds. are

Searcher :

Shears

09/787436

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further provided, as are methods for preparing a radiotherapeutic agent from
     the compds.
     ICM A61K
IC
     63-5 (Pharmaceuticals)
     Section cross-reference(s): 1, 8, 26, 34
     imaging agent targeted conjugate prepn bile acid; radiotherapeutic
     targeted conjugate prepn
IT
     Imaging agents
        (NMR contrast; targeted diagnostic and therapeutic agents with improved
        half life)
IT
     Imaging agents
        (acoustic; targeted diagnostic and therapeutic agents with improved
        half life)
IT
     Antibiotics
        (conjugates; targeted diagnostic and therapeutic agents with improved
IΤ
     Antibodies and Immunoglobulins
     Interleukin 1
     Neurokinins
     RL: DGN (Diagnostic use); THU (Therapeutic use); BIOL (Biological study);
        (conjugates; targeted diagnostic and therapeutic agents with improved
        half life)
IT
     Enzymes, biological studies
     Growth factors, animal
     Hormones, animal, biological studies
     RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)
        (conjugates; targeted diagnostic and therapeutic agents with improved
        half life)
ΙT
     Drug delivery systems
        (immunoconjugates; targeted diagnostic and therapeutic agents with
        improved half life)
IT
     Drug delivery systems
        (injections; targeted diagnostic and therapeutic agents with improved
        half life)
IT
     Human
     Imaging agents
     Photodynamic therapy
     Phototherapy
     Radiopharmaceuticals
     Radiotherapy
     Test kits
        (targeted diagnostic and therapeutic agents with improved half life)
ΙT
     Amino acids, biological studies
     Bile acids
     RL: DGN (Diagnostic use); THU (Therapeutic use); BIOL (Biological study);
     USES (Uses)
        (targeted diagnostic and therapeutic agents with improved half life)
     7440-54-2D, Gadolinium, cholic acid and peptide-conjugated complexes
IT
     RL: DGN (Diagnostic use); BIOL (Biological study); USES (Uses)
        (targeted diagnostic and therapeutic agents with improved half life)
IT
     721937-44-6DP, Lutetium 177 complexes
                                            721937-46-8DP, Lutetium 177/Indium
     111/Gadolinium complexes
     RL: DGN (Diagnostic use); PAC (Pharmacological activity); PKT
     (Pharmacokinetics); SPN (Synthetic preparation); THU (Therapeutic use);
     BIOL (Biological study); PREP (Preparation); USES (Uses)
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(targeted diagnostic and therapeutic agents with improved half life)
IT
     14265-75-9DP, 177Lu, cholic acid and peptide-conjugated complexes,
     biological studies
     RL: DGN (Diagnostic use); PAC (Pharmacological activity); SPN (Synthetic
     preparation); THU (Therapeutic use); BIOL (Biological study); PREP
     (Preparation); USES (Uses)
         (targeted diagnostic and therapeutic agents with improved half life)
IT
     83-44-3DP, Deoxycholic acid, conjugates 14158-31-7DP, Iodine 125,
     insulin derivs. labeled with, biological studies
                                                        721937-42-4DP,
     complexes
                 721937-44-6DP, complexes
                                           721937-46-8DP, complexes
     721937-48-0DP, complexes
                                721937-50-4DP, complexes
                                                           721937-52-6DP.
                 721937-54-8DP, complexes
     complexes
                                           728038-66-2DP, complexes
     728919-39-9DP, Indium 111 complexes
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     use); BIOL (Biological study); PREP (Preparation); USES (Uses)
        (targeted diagnostic and therapeutic agents with improved half life)
     50-56-6D, Oxytocin, conjugates
                                     58-82-2D, Bradykinin, conjugates
     69-79-4D, Maltose, conjugates
                                    113-79-1D, Arginine vasopressin,
                  9002-79-3D, MSH, conjugates 9011-97-6D, CCK, conjugates
     conjugates
                                     15750-15-9D, Indium 111, conjugated
     9034-40-6D, LH-RH, conjugates
     complexes, biological studies
                                     33507-63-0D, Substance P, conjugates
     37221-79-7D, VIP, conjugates
                                    51110-01-1D, Somatostatin, conjugates
     62229-50-9D, EGF, conjugates
                                    82785-45-3D, Neuropeptide Y, conjugates
     83150-76-9D, Octreotide, conjugates
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                                           116243-73-3D, Endothelin,
     108736-35-2D, Lanreotide, conjugates
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             108-30-5, reactions 29022-11-5 38359-38-5
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        (targeted diagnostic and therapeutic agents with improved half life)
IT
     9004-10-8D, Insulin, conjugates
    RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)
        (targeted diagnostic and therapeutic agents with improved half life)
L36 ANSWER 2 OF 6 MARPAT COPYRIGHT 2004 ACS on STN
ACCESSION NUMBER:
                         141:47383 MARPAT
TITLE:
                         Peptides having antiangiogenic activity
INVENTOR(S):
                         Haviv, Fortuna; Henkin, Jack; Bradley, Michael F.;
                         Kalvin, Douglas M.; Schneider, Andrew J.
PATENT ASSIGNEE(S):
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SOURCE:
                         U.S., 27 pp.
                         CODEN: USXXAM
DOCUMENT TYPE:
                         Patent
LANGUAGE:
                         English
FAMILY ACC. NUM. COUNT:
PATENT INFORMATION:
     PATENT NO.
                      KIND DATE
                                          APPLICATION NO. DATE
                                           _____
    US 6753408
                                           US 2000-718591
                      В1
                            20040622
                                                            20001122
PRIORITY APPLN. INFO.:
                                           US 1999-166791P 19991122
     Peptides Xaa1-Xaa2-Xaa3-Xaa4-Xaa5-Xaa6-Xaa7-Xaa8-Xaa9-Xaa10-Xaa11 are
     useful for inhibiting angiogenesis. Also disclosed are
     angiogenesis-inhibiting compns. and methods of inhibiting angiogenesis in
     a mammal. Compds. such as N-Ac-Sar-Gly-Lys(Ac)-D-Leu-Thr-Nva-Ile-Arg-
     ProNH-Et were prepared The compds. inhibited human endothelial cell
     migration by at least 50 % inhibition when tested at concns. of 1 nM.
IC
     ICM C07K007-00
NCL
    530328000
CC
     1-12 (Pharmacology)
     Section cross-reference(s): 34
ST
     antiangiogenic peptide; endothelial cell migration inhibition peptide
ΙT
        (microvessel, endothelium, inhibition of migration of cells of, of
        human; peptides having antiangiogenic activity)
IT
     Cell migration
        (of human microvascular endothelial cells, inhibition of; peptides
        having antiangiogenic activity)
IT
     Angiogenesis inhibitors
     Antitumor agents
     Drug delivery systems
     Human
     Mammalia
        (peptides having antiangiogenic activity)
TT
     Peptides, biological studies
     RL: BSU (Biological study, unclassified); PAC (Pharmacological activity);
     THU (Therapeutic use); BIOL (Biological study); USES (Uses)
        (peptides having antiangiogenic activity)
                    341522-18-7P 341522-20-1P
IT
     341522-16-5P
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        (peptides having antiangiogenic activity)
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IT

341522-17-6P

Searcher : Shears 571-272-2528

341522-23-4P

341522-25-6P

341522-19-8P 341522-21-2P

341522-34-7P

341522-36-9P

341522-38-1P

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المارين يومي

341522-30-3P

341522-32-5P

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341522-40-5P
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     RL: PAC (Pharmacological activity); PRP (Properties); SPN (Synthetic
     preparation); THU (Therapeutic use); BIOL (Biological study); PREP
     (Preparation); USES (Uses)
        (peptides having antiangiogenic activity)
                               THERE ARE 24 CITED REFERENCES AVAILABLE FOR THIS
REFERENCE COUNT:
                         24
                               RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT
L36 ANSWER 3 OF 6 MARPAT COPYRIGHT 2004 ACS on STN
                         140:321722 MARPAT
ACCESSION NUMBER:
TITLE:
                         Preparation of peptide antiangiogenic drugs
INVENTOR(S):
                         Henkin, Jack; Haviv, Fortuna; Bradley, Michael F.;
                         Kalvin, Douglas M.; Schneider, Andrew J.
PATENT ASSIGNEE(S):
                         Abbott Laboratories, USA
SOURCE:
                         U.S., 84 pp., Cont.-in-part of U.S. Ser. No. 316,888.
                         CODEN: USXXAM
DOCUMENT TYPE:
                         Patent
LANGUAGE:
                         English
FAMILY ACC. NUM. COUNT:
PATENT INFORMATION:
     PATENT NO.
                      KIND DATE
                                           APPLICATION NO.
                                                            DATE
     ____
    US 6716963
                                           US 1999-447226
                       В1
                            20040406
                                                            19991122
    US 6774211
                       В1
                            20040810
                                           US 2001-833196
                                                            20010411
PRIORITY APPLN. INFO.:
                                           US 1998-86536P
                                                            19980522
                                           US 1999-126546P 19990326
                                           US 1999-316888
                                                            19990521
                                           US 1999-447226
                                                            19991122
AΒ
     Peptides A0-A1-A2-A3-A4-A5-A6-A7-A8-A9-A10 (A0 is an acyl group; A10 is
     OH, an amino group, or an amino acid amide; A1-9 are amino acyl residues)
     or their pharmaceutically acceptable salts, esters, solvates, or prodrugs
     were prepared for the treatment of angiogenesis. Thus, N-Ac-Sar-Gly-Val-D-
     Ile-Thr-Nva-Ile-Arg-Pro-NHEt was prepared by the solid-phase method and
     assayed for in vitro angiogenic activity (87.3 and 76.9% inhibition at 20
     nM and 10 nM. resp.).
     ICM C07K007-00
IC
    530328000
NCL
     34-3 (Amino Acids, Peptides, and Proteins)
     Section cross-reference(s): 1
ST
     peptide prepn antiangiogenic activity
IT
     Solid phase synthesis
        (peptide; preparation of peptide antiangiogenic drugs)
IT
    Angiogenesis
     Angiogenesis inhibitors
    Antitumor agents
    Neoplasm
        (preparation of peptide antiangiogenic drugs)
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EX. 9. 1.

Peptides, preparation

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RL: PAC (Pharmacological activity); SPN (Synthetic preparation); THU
     (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES
     (Uses)
        (preparation of peptide antiangiogenic drugs)
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    RL: PAC (Pharmacological activity); SPN (Synthetic preparation); THU
    (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES
    (Uses)
        (preparation of peptide antiangiogenic drugs)
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     RL: PAC (Pharmacological activity); SPN (Synthetic preparation); THU
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        (preparation of peptide antiangiogenic drugs)
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(Therapeutic use); BIOL (Biological study); PREP (Preparation); USES
(Uses)
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(preparation of peptide antiangiogenic drugs)

TT 59-67-6, Nicotinic acid, reactions 79-09-4, Propionic acid, reactions 88-14-2, 2-Furoic acid 107-92-6, Butanoic acid, reactions 109-85-3, 2-Methoxyethylamine 138-59-0, Shikimic acid 142-62-1, Hexanoic acid, reactions 625-45-6, Methoxyacetic acid 2516-47-4,

09/787436

3222-56-8, 2-Methylnicotinic acid 4442-85-7, Cyclopropanemethanamine 5292-21-7, Cyclohexylacetic acid 591: 7154-73-6, 1-(2-Aminoethyl)pyrrolidine 2-Cyclohexylethylamine 5913-13-3, r 1-Cyclohexylethylamine 16874-33-2, Tetrahydro-2-furoic acid 27578-60-5, 1-(2-Aminoethyl)piperidine 101711-55-1 RL: RCT (Reactant); RACT (Reactant or reagent) (preparation of peptide antiangiogenic drugs) IT 675886-06-3 675886-07-4 675886-08-5 675886-09-6 RL: PRP (Properties) (unclaimed protein sequence; preparation of peptide antiangiogenic drugs) 30 REFERENCE COUNT: THERE ARE 30 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT L36 ANSWER 4 OF 6 MARPAT COPYRIGHT 2004 ACS on STN ACCESSION NUMBER: 135:5821 MARPAT TITLE: Preparation of peptides having antiangiogenic activity INVENTOR(S): Haviv, Fortuna; Henkin, Jack; Bradley, Michael F.; Kalvin, Douglas M.; Schneider, Andrew J. PATENT ASSIGNEE(S): Abbott Laboratories, USA SOURCE: PCT Int. Appl., 79 pp. CODEN: PIXXD2 DOCUMENT TYPE: Patent English LANGUAGE: FAMILY ACC. NUM. COUNT: PATENT INFORMATION: PATENT NO. KIND DATE APPLICATION NO. ____ -----WO 2001038347 A2 20010531 WO 2000-US32217 20001122 WO 2001038347 Α3 20011129 W: CA, JP, MX RW: AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, TR CA 2000-2391386 20001122 CA 2391386 20010531 AΑ EP 1232183 20020821 Α2 EP 2000-982219 20001122 R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, FI, CY, TR JP 2003530313 JP 2001-540110 T2 20031014 20001122 PRIORITY APPLN. INFO.: US 1999-447225 19991122 US 2000-709034 20001108 WO 2000-US32217 20001122 AΒ Peptides Xaa1-Xaa2-Xaa3-Xaa4-Xaa5-Xaa6-Xaa7-Xaa8-Xaa9-Xaa10-Xaa11 [Xaa1 is absent, H or an acyl group; Xaa2-Xaa10 represent certain amino acyl groups; Xaall is OH, an amino acid amide selected from D-alanylamide, D-alanylethylamide, azaglycylamide, glycylamide, glycylethylamide, sarcosylamide, serylamide, D-serylamide, a residue NH(CH2)sCHR3R4 or NHR5 [s = 0-8, R3 = H, alkyl, 5- to 6-membered cycloalkyl; R4 = H, alkoxy, alkyl, aryl, cycloalkenyl, cycloalkyl, heterocyclyl, OH; R5 = H, OH, cycloalkyl (with provisos)]] or their pharmaceutically acceptable salts were prepared for inhibiting angiogenesis. Thus, Ac-Sar-Gly-Lys(Ac)-D-Leu-Thr-Nva-Ile-Arg-Pro-NHEt was prepared by the solid-phase method using Fmoc-protected amino acids. The synthesized peptides inhibited human

34-3 (Amino Acids, Peptides, and Proteins) Section cross-reference(s): 1

endothelial cell migration by at least 50 % at concns. of 1 nM.

200 to 100 to

Sec. 2.

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CC

ICM C07K007-00

Searcher : Shears

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         (diabetic retinopathy; preparation of peptides having antiangiogenic
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     Eye, disease
        (macula, degeneration; preparation of peptides having antiangiogenic
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IT
     Angiogenesis inhibitors
     Antiarthritics
     Antitumor agents
     Psoriasis
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        (preparation of peptides having antiangiogenic activity)
L36 ANSWER 5 OF 6 MARPAT COPYRIGHT 2004 ACS on STN
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                         Preparation of peptide antiangiogenic drugs
INVENTOR(S):
                         Henkin, Jack; Haviv, Fortuna; Bradley, Michael F.;
                         Kalvin, Douglas M.; Schneider, Andrew J.
PATENT ASSIGNEE(S):
                         Abbott Laboratories, USA
SOURCE:
                         PCT Int. Appl., 223 pp.
                         CODEN: PIXXD2
DOCUMENT TYPE:
                         Patent
LANGUAGE:
                         English
FAMILY ACC. NUM. COUNT:
PATENT INFORMATION:
     PATENT NO.
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Searcher : 571-272-2528 Shears

APPLICATION NO.

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     is OH or an amino acid amide; Al-9 are amino acyl residues) or their
     pharmaceutically acceptable salts, esters, solvates, or prodrugs were
     prepared for the treatment of angiogenesis. Thus, N-Ac-Sar-Gly-Val-D-Ile-
     Thr-Nva-Ile-Arg-Pro-NHEt was prepared by the solid-phase method and assayed
     for in vitro angiogenic activity (87.3% at 20 nM and 76.9 at 10 nM).
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     Eye, disease
        (diabetic retinopathy; preparation of peptide antiangiogenic drugs)
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        (macula, degeneration; preparation of peptide antiangiogenic drugs)
     Angiogenesis inhibitors
     Antiarthritics
     Antitumor agents
     Psoriasis
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     59-67-6, Nicotinic acid, reactions 79-09-4, Propionic acid, reactions
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     88-14-2, 2-Furoic acid
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     1-Cyclohexylethylamine
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REFERENCE COUNT:
                               THERE ARE 4 CITED REFERENCES AVAILABLE FOR THIS
                               RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT
L36 ANSWER 6 OF 6 MARPAT COPYRIGHT 2004 ACS on STN
ACCESSION NUMBER:
                         125:87217 MARPAT
TITLE:
                         Preparation of polymer-bound luteinizing hormone
                         releasing factor analogs with antitumor activity.
INVENTOR(S):
                         Lovas, Sandor; Murphy, Richard F.; Toth, Geza; Kalnay,
                         Adrienn; Gaal, Dezso; Palyi, Istvan; Turi, Gizella;
                         Vincze, Borbala; Mezo, Imre; et al.
PATENT ASSIGNEE(S):
                         PCT Int. Appl., 70 pp.
SOURCE:
                         CODEN: PIXXD2
DOCUMENT TYPE:
                         Patent
LANGUAGE:
                         English
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FAMILY ACC. NUM. COUNT: 1 PATENT INFORMATION:

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PRIORITY APPLN. INFO.:
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                                                            19940810
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     the other = 2-oxopyrrolidin-1-yl; R3 = polymerization-initiating group,
     preferably Me2CCN; W = OH (or alkali metal salt thereof); V = alkylamino,
     bond; X = amino acid, r oligopeptide; A = pharmacol. active polypeptide; r
     = 0 to 0.2n; k \le r; z = 0 to n-r; u = n to 2n-r] and
     nonpolymer-bound peptides, were prepared Thus, N-vinylpyrrolidone-maleic
     anhydride copolymer was stirred 3 h with H-Gly-Phe-Leu-Gly nitrophenyl
     ester and Et3N in DMF to give after aqueous hydrolysis
poly(N-Vinylpyrrolidone-
     maleic acid)-Gly-Phe-Leu-Gly nitrophenyl ester. The latter in DMF was
     stirred 24 h with Ac-D-Trp1, 3, D-Cpa2, Lys5, [β-Asp(DEA)]6, D-Ala10-HGnRH
     and Et3N to give Ac-D-Trp1,3,D-Cpa2,Lys[poly(N-vinylpyrrolidone-maleic
     acid)-Gly-Phe-Leu-Gly-] 5, [\beta-Asp(DEA)]6, D-Ala10-HGnRH. The latter at
     50 µM in MCF-7 cell cultures showed 93% inhibition of colony formation.
IC
     ICM A61K038-00
     ICS C07K005-00; C07K007-00; C07K017-00
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ST
     gnrh analog polymer bound prepn anticancer
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     Immunostimulants
     Neoplasm inhibitors
        (preparation of polymer-bound GNRH analogs with antitumor activity)
IT
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     study, unclassified); SPN (Synthetic preparation); THU (Therapeutic use);
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Searcher: Shears 571-272-2528

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26837-56-9, N-Vinylpyrrolidone-maleic anhydride copolymer 52671-12-2
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147859-97-0, Luteinizing hormone-releasing factor III (Petromyzon marinus)
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(preparation of polymer-bound GNRH analogs with antitumor activity)

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NODE ATTRIBUTES: DEFAULT MLEVEL IS ATOM MLEVEL IS CLASS AT 16 29 GGCAT IS PCY AT 16

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GRAPH ATTRIBUTES:

RING(S) ARE ISOLATED OR EMBEDDED

NUMBER OF NODES IS 36

STEREO ATTRIBUTES: NONE

ATTRIBUTES SPECIFIED AT SEARCH-TIME:

ECLEVEL IS LIM ON ALL NODES

ALL RING(S) ARE ISOLATED

L37 0 SEA FILE=MARPATPREV SSS FUL L32 (MODIFIED ATTRIBUTES)

100.0% PROCESSED 12 ITERATIONS

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SEARCH TIME: 00.00.01

FILE 'REGISTRY' ENTERED AT 13:01:07 ON 22 NOV 2004

L38 4 S EHWSYL/SQSP

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L38 ANSWER 1 OF 4 REGISTRY COPYRIGHT 2004 ACS on STN

RN 581927-52-8 REGISTRY

CN Protein (Klebsiella pneumoniae strain ATCC202080 clone

US6610836-SEQID-13292 open reading frame-encoded) (9CI) (CA INDEX NAME)

OTHER NAMES:

CN 2592: PN: US6610836 SEQID: 13292 claimed protein

CI MAN

Sec. 4.

SQL 573

SEQ 1 SPSRLRWVNS SPYFNREVFL QFDYIIIGAG SAGNVLATRL TEDPNTTVLL

51 LEAGGPDYRF DFRTQMPAAL AYPLQGKRYN WAYETEPEPY MNNRRMECGR

101 GKGLGGSSLI NGMCYIRGNA MDLDNWAKEP GLEHWSYLDC LPYYRKAETR

151 DIGPNDYHGG DGPVSVTTPK PGNNPLFEAM VEAGVQAGYP RTDDLNGYQQ

201 EGFGPMDRTV TPQGRRASTA RGYLDQARGR PNLTIRTHAL TDHIIFAGKR

251 AVGVEWLEGE STIPSKATAN KEVLLCAGAI ASPQILQRSG VGNPELLRQF

301 DIPVVHDLPG VGENLQDHLE MYLQYECKEP VSLYPALQWW NQPKIGAEWL

351 FGGTGIGASN QFEAGGFIRS RAEFAWPNIQ YHFLPVAINY NGSNAVKEHG

401 FQCHVGSMRS PSRGHVRLKS RDPHAHPAIL FNYMSHEQDW QEFRDAIRIT

451 REIMNQPALD KYRGREISPG IECQSDAELD EFVRNHAETA FHPCGTCKMG

501 YDEMAVVDGE GRVHGLEGLR VVDASIMPQI ITGNLNATTI MIGEKMADAI

551 RGRQPLPRST ATYYVAGDAP VRR

HITS AT: 133-138

REFERENCE 1: 139:192571

L38 ANSWER 2 OF 4 REGISTRY COPYRIGHT 2004 ACS on STN

RN 477074-93-4 REGISTRY

CN Protein (Klebsiella pneumoniae clone KPN304388 essential) (9CI) (CA INDEX NAME)

OTHER NAMES:

CN 2036: PN: W002077183 SEQID: 60036 claimed protein

CI MAN

SQL 554

22 P. S. S.

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Service 1

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SEQ
         1 MOFDYIIIGA GSAGNVLATR LTEDPNTTVL LLEAGGPDYR FDFRTOMPAA
        51 LAYPLOGKRY NWAYETEPEP YMNNRRMECG RGKGLGGSSL INGMCYIRGN
       101 AMDLDNWAKE PGLEHWSYLD CLPYYRKAET RDIGPNDYHG GDGPVSVTTP
       151 KPGNNPLFEA MVEAGVQAGY PRTDDLNGYQ QEGFGPMDRT VTPQGRRAST
       201 ARGYLDOARG RPNLTIRTHA LTDHIIFAGK RAVGVEWLEG ESTIPSKATA
       251 NKEVLLCAGA IASPQILQRS GVGNPELLRQ FDIPVVHDLP GVGENLQDHL
       301 EMYLQYECKE PVSLYPALQW WNQPKIGAEW LFGGTGIGAS NQFEAGGFIR
       351 SRAEFAWPNI QYHFLPVAIN YNGSNAVKEH GFQCHVGSMR SPSRGHVRLK
       401 SRDPHAHPAI LFNYMSHEQD WQEFRDAIRI TREIMNQPAL DKYRGREISP
       451 GIECOSDAEL DEFVRNHAET AFHPCGTCKM GYDEMAVVDG EGRVHGLEGL
       501 RVVDASIMPQ IITGNLNATT IMIGEKMADA IRGRQPLPRS TATYYVAGDA
       551 PVRR
           114-119
HITS AT:
REFERENCE
            1: 138:1095
     ANSWER 3 OF 4 REGISTRY COPYRIGHT 2004 ACS on STN
     465609-93-2 REGISTRY
     Protein (Plasmodium falciparum strain 3D7 gene PF10-0219) (9CI)
                                                                       (CA INDEX
     NAME)
OTHER NAMES:
CN
     GenBank AAN35416
     GenBank AAN35416 (Translated from: GenBank AE014832)
CN
CI
     MAN
SQL
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         1 MVKKDKKSKE KKEKLKLKKE KQKLKSLKSK KKKKDTLSDE DFDTICLYYE
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       101 NDNNELISYN DLFKYNIVKN KWKYYFTTSK KPKPRCSHQT VYFNKKLYIF
       151 GGELCTNTQF FHYNDFWSFD LKNNVFEEIE TKNKKDDNKP SPRSGHRMIL
       201 WKNYIVMFGG FFDNGKSIEY FNDLYIYIIN SNIWINLTNV YMDSLFKRLT
       251 ENNSSNNDNN LSISSEKKKD LNKGKNSQIL KSKFFKNFDL DSFMPSKRSS
       301 VCLFTDMKYQ KIYIYGGYSQ IKNTTRNAIG FYFNDMWILN INLINEDNIS
       351 VNFKKLKKSI FQPCKRTGFS TCIYKNSLIL FGGVFDKKVE NNSKKIDNPN
       401 NNNNMLEESL NLKSLFFNDL YLFDMNKEHW SYLNIKDKEE TKELNKTSKA
       451 NKKNHEKLEE NKIGTNIKKD KFQQMEREIY YEENNNNNNN NNNNKTQSKY
       501 EETSDGHVSS CFSDDNDEDY YSNVFVYFDE NGKRQIIKIE KEEKNKSSYN
       551 EKKDFDDVLK VEENNDYLNH SLDEEKNINI DKLIDNHSVF LQTKDIITTG
       601 NGNKVTKIFG DENKHCQNIS NLPLNETILY VPLNNTTNSE NFMYSQELND
       651 STNMIKVEHI NDTENVDEET CKEDSVDEDM KDNSNSDSDS NKEEKKKKFV
       701 ISEEEPIGRI NSHIFVLNKN LYVYGGMYEY KNNEIILSDY WKINIFKREK
       751 WELLDKGNLD DIYLEESDMS STISINDDDK DEKEIEDLII CSKIKKLEKK
       801 IKELDEGLAL DIKENLNEFF LRTKDHWLKE LNKISETKEI RKEAFYLCEQ
       851 KYKVIKKYYN KIQKYKELLM EDDEERSISE TISSEQEQSS N
HITS AT:
           428-433
REFERENCE
            1: 137:289734
    ANSWER 4 OF 4 REGISTRY COPYRIGHT 2004 ACS on STN
RN
     112642-13-4 REGISTRY
     Luteinizing hormone-releasing factor (swine), 1-L-glutamic
     acid-6-D-leucine-9-(N-ethyl-L-prolinamide)-10-deglycinamide- (9CI)
     INDEX NAME)
OTHER CA INDEX NAMES:
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CN Luteinizing hormone-releasing factor (pig), 1-L-glutamic acid-6-D-leucine-9-(N-ethyl-L-prolinamide)-10-deglycinamide-

SQL 9

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-212 ·

SEQ 1 EHWSYLLRP

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HITS AT: 1-6

REFERENCE 1: 108:82207

FILE 'CAPLUS' ENTERED AT 13:01:52 ON 22 NOV 2004

L39 4 S L38

L40 3 S L39 NOT L28

L40 ANSWER 1 OF 3 CAPLUS COPYRIGHT 2004 ACS on STN

ED Entered STN: 07 Sep 2003

ACCESSION NUMBER: 2003:697220 CAPLUS

DOCUMENT NUMBER: 139:192571

TITLE: Nucleic acid and encoded amino acid sequences relating

to Klebsiella pneumoniae for diagnostics and

APPLICATION NO.

DATE

therapeutics

INVENTOR(S): Breton, Gary L.; Osborne, Mark

KIND

PATENT ASSIGNEE(S): Genome Therapeutics Corporation, USA

SOURCE: U.S., 932 pp.

CODEN: USXXAM

DATE

DOCUMENT TYPE: Patent LANGUAGE: English

FAMILY ACC. NUM. COUNT: 4

PATENT INFORMATION:

PATENT NO.

	US 6610836	B1	20030826	US 2000-489039		20000127			
PRIO	RITY APPLN. INFO.:			US 1999-117747P	Р	19990129			
AB	The invention provi	ides 717	71 isolated	polypeptide and 717	l gen	omic nucleic			
	acid sequences derived from Klebsiella pneumoniae strain 93,19097 (ATCC								
202080) that are useful in diagnosis and therapy of pathol. conditions.									
The nucleotide sequences include those of two naturally occurring plasmids									
in K. pneumoniae. Antibodies against the polypeptides, and methods for									
the production of recombinant polypeptides are also provided. The invention									
also provides methods for the detection, prevention, and treatment of									
pathol. conditions resulting from bacterial infection. [This abstract									
	record is one of four records for this document necessitated by the large								
	number of index ent	ries re	equired to f	ully index the docu	nent	and publication			
	system constraints.		•			F			

IT 581927-52-8

RL: BSU (Biological study, unclassified); DGN (Diagnostic use); PRP (Properties); THU (Therapeutic use); BIOL (Biological study); USES (Uses) (amino acid sequence; nucleic acid and encoded amino acid sequences relating to Klebsiella pneumoniae for diagnostics and therapeutics)

L40 ANSWER 2 OF 3 CAPLUS COPYRIGHT 2004 ACS on STN

ED Entered STN: 14 Oct 2002

ACCESSION NUMBER: 2002:781491 CAPLUS

DOCUMENT NUMBER: 138:1095

TITLE: Essential genes in microorganisms and their use as

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targets for antisense inhibition of proliferation and
                         antibiotic screening
INVENTOR(S):
                        Wang, Liangus; Zamudio, Carlos; Malone, Cheryl;
                        Haselbeck, Robert; Ohlsen, Kari L.; Zyskind, Judith
                        W.; Wall, Daniel; Trawick, John D.; Carr, Grant J.;
                        Yamamoto, Robert; Forsyth, R. Allyn; Xu, H. Howard
PATENT ASSIGNEE(S):
                        Elitra Pharmaceuticals, Inc., USA
SOURCE:
                        PCT Int. Appl., 1766 pp.
                        CODEN: PIXXD2
DOCUMENT TYPE:
                        Patent
LANGUAGE:
                        English
FAMILY ACC. NUM. COUNT:
                        22
PATENT INFORMATION:
     PATENT NO.
                        KIND
                               DATE
                                           APPLICATION NO.
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WO 2002077183
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                CO, CR, CU, CZ, DE, DE, DK, DK, DM, DZ, EC, EE, EE, ES, FI, FI,
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                LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO,
                NZ, OM, PH, PL, PT, RO, RU, SD, SE, SG, SI, SK, SK, SL, TJ, TM,
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      US 2002061569
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                                         20020523
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                                                                                     20010321
      WO 2002077183
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                                 A2
                                                                                     20020321
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                BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG
PRIORITY APPLN. INFO.:
                                                        US 2001-815242
                                                                               A 20010321
                                                        US 2001-948993
                                                                                 A 20010906
                                                        US 2001-342923P
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                                                        US 2002-72851
                                                                                 A 20020208
                                                        US 2002-362699P
                                                                                 P 20020306
                                                        WO 2002-US9107
                                                                                 A 20020321
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                                                        US 2000-257931P
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                                                                                     20001222
                                                        US 2001-269308P
                                                                                 Р
                                                                                     20010216
      The sequences of antisense nucleic acids which inhibit the proliferation
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AB The sequences of antisense nucleic acids which inhibit the proliferation of prokaryotes are disclosed. Thus, 6213 nucleic acid fragments are identified for which expression inhibits proliferation or is required for proliferation in Enterococcus faecalis, Escherichia coli, Klebsiella pneumoniae, Pseudomonas aeruginosa, Salmonella typhimurium, and

Staphylococcus aureus. Cell-based assays which employ the antisense nucleic acids to identify and develop antibiotics are also disclosed. antisense nucleic acids can also be used to identify proteins required for proliferation, express these proteins or portions thereof, obtain antibodies capable of specifically binding to the expressed proteins, and to use those expressed proteins as a screen to isolate candidate mols. for rational drug discovery programs. The nucleic acids can also be used to screen for homologous nucleic acids that are required for proliferation in cells other than Staphylococcus aureus, Salmonella typhimurium, Klebsiella pneumoniae, and Pseudomonas aeruginosa. The invention provides 38,184 such proliferation-required gene sequences (plus their encoded protein sequences). The nucleic acids of the present invention can also be used in various assay systems to screen for proliferation required genes in other organisms. [This abstract record is one of twenty records for this document necessitated by the large number of index entries required to fully index the document and publication system constraints.].

IT

RL: BSU (Biological study, unclassified); BUU (Biological use, unclassified); PRP (Properties); THU (Therapeutic use); BIOL (Biological study); USES (Uses)

(amino acid sequence; essential genes in microorganisms and their use as targets for antisense inhibition of proliferation and antibiotic screening)

ANSWER 3 OF 3 CAPLUS COPYRIGHT 2004 ACS on STN

04 Oct 2002 Entered STN:

ACCESSION NUMBER: 2002:752115 CAPLUS

DOCUMENT NUMBER:

137:289734

TITLE:

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Sequence of Plasmodium falciparum chromosomes 2, 10,

AUTHOR(S):

Gardner, Malcolm J.; Shallom, Shamira J.; Carlton, Jane M.; Salzberg, Steven L.; Nene, Vishvanath; Shoaibi, Azadeh; Ciecko, Anne; Lynn, Jeffery; Rizzo, Michael; Weaver, Bruce; Jarrahi, Behnam; Brenner, Michael; Parvizi, Babak; Tallon, Luke; Moazzez, Azita; Granger, David; Fujii, Claire; Hansen, Cheryl; Pederson, James; Feldblyum, Tamara; Peterson, Jeremy; Suh, Bernard; Angiuoli, Sam; Pertea, Mihaela; Allen, Jonathan; Selengut, Jeremy; White, Owen; Cummings, Leda M.; Smith, Hamilton O.; Adams, Mark D.; Venter, J. Craig; Carucci, Daniel J.; Hoffman, Stephen L.; Fraser, Claire M.

CORPORATE SOURCE:

The Institute for Genomic Research, Rockville, MD,

20850, USA

SOURCE:

Nature (London, United Kingdom) (2002), 419(6906),

531-534

CODEN: NATUAS; ISSN: 0028-0836

PUBLISHER:

DOCUMENT TYPE:

Nature Publishing Group Journal

English

LANGUAGE:

The mosquito-borne malaria parasite Plasmodium falciparum kills an estimated 0.7-2.7 million people every year, primarily children in sub-Saharan Africa. Without effective interventions, a variety of factors-including

the spread of parasites resistant to antimalarial drugs and the increasing insecticide resistance of mosquitoes-may cause the number of malaria cases

to

Searcher :

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double over the next two decades. To stimulate basic research and facilitate the development of new drugs and vaccines, the genome of Plasmodium falciparum clone 3D7 has been sequenced using a chromosome-by-chromosome shotgun strategy. This report describes nucleotide sequences of chromosomes 10, 11 and 14, and a re-anal. of the chromosome 2 sequence. These chromosomes represent about 35% of the 23-megabase P. falciparum genome. The sequences are deposited in GenBank/EMBL/DDBJ under accession nos. AE001362.2 (chromosome 2), AE014185 (chromosome 10), AE014186 (chromosome 11), and AE014187 (chromosome 14).

IT 465609-93-2

25 P. F. S.

RL: BSU (Biological study, unclassified); PRP (Properties); BIOL (Biological study)

(amino acid sequence; complete sequence of Plasmodium falciparum chromosomes 2, 10, 11 and 14)

REFERENCE COUNT:

THERE ARE 30 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

(FILE 'MEDLINE, BIOSIS, EMBASE' ENTERED AT 13:02:33 ON 22 NOV 2004) L41 0 S L38

FILE 'HOME' ENTERED AT 13:02:41 ON 22 NOV 2004

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